

- <110> Oppermann, Herman Kuberasampath, Thangavel Rueger, David Ozkaynak, Engin
- <120> Osteogenic Devices
- <130> STK-008CN
- <140> US 09/754,831
- <141> 2001-01-03
- <150> US 08/375,901
- <151> 1995-01-20
- <150> US 08/145,812
- <151> 1993-11-01
- <150> US 07/995,345
- <151> 1992-12-22
- <150> US 07/315,342
- <151> 1989-02-23
- <150> US 07/232,630
- <151> 1988-08-15
- <150> US 07/179,406
- <151> 1988-04-08
- <160> 72
- <170> PatentIn version 3.0
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- <212> PRT
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- Val Asn Ser Val Asn Ser Lys Ile Pro Lys Ala Cys Cys Val Pro Thr 50 55 60

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                         . 40
Val Asn Ser Val Asn Ser Lys Ile Pro Lys Ala Cys Cys Val Pro Thr
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            2.0
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Asn His Ala Xaa Xaa Gln Xaa Xaa
Val Xaa Xaa Xaa Asn Xaa Xaa Xaa Pro Xaa Xaa Cys Cys Xaa Pro
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Leu Xaa Xaa Xaa Xaa Xaa Xaa
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Xaa
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Xaa Cys Xaa Xaa Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Asn His Ala
       35
                           40
                                               45
Xaa Xaa Gln Xaa Xaa Val Xaa Xaa Xaa Asn Xaa Xaa Xaa Pro Xaa
Xaa Cys Cys Xaa Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Leu Xaa Xaa
                    70
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<223> wherein Xaa at position 11 is an asparagine, a glutamine, an
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<223> wherein Xaa at position 23 is a phenylalanine, a tyrosine or an
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      (26)..(26)
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      (31)..(31)
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Val Xaa Xaa Xaa Asn Xaa Xaa Xaa Pro Xaa Xaa Cys Cys Xaa Pro
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Leu Xaa Xaa Xaa Xaa Xaa Xaa
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<223> wherein Xaa at position 25 is a tyrosine or a phenylalanine
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<222> (44)..(44)
<223> wherein Xaa at position 44 is a serine, an alanine or a proline
<220>
<221> misc_feature
<222> (45)..(45)
<223> wherein Xaa at position 45 is a threonine or a serine
<220>
<221> misc_feature
<222> (49)..(49)
<223> wherein Xaa at position 49 is an isoleucine, a valine or a threonine
<220>
<221> misc feature
<222> (50)..(50)
<223> wherein Xaa at position 50 is a valine, an isoleucine or a leucine
<220>
<221> misc feature
<222> (52)..(52)
<223> wherein Xaa at position 52 is a threonine or a serine
<220>
<221> misc_feature
<222> (53)..(53)
<223> wherein Xaa at position 53 is a leucine or a isoleucine
<220>
<221> misc_feature
<222> (55)..(55)
<223> wherein Xaa at position 55 is an asparagine, a histidine or an
      arginine
<220>
<221> misc_feature
<222> (56)..(56)
<223> wherein Xaa at position 56 is a serine, an alanine, a phenylalanine
      or an asparagine
<220>
<221> misc feature
<222> (57)..(57)
<223> wherein Xaa at position 57 is a valine or an isoleucine
<220>
<221> misc feature
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<222> (59)..(59)
<223> wherein Xaa at position 59 is a serine or a proline
<220>
<221> misc feature
<222> (60)..(60)
<223> wherein Xaa at position 60 is a glycine or a glutamic acid
<220>
<221> misc feature
<222> (61)..(61)
<223> wherein Xaa at position 61 is a lysine, a glutamine, a threonine
      or a serine
<220>
<221> misc_feature
<222> (62)..(62)
<223> wherein Xaa at position 62 is an isoleucine or a valine
<220>
<221> misc feature
<222> (64)..(64)
<223> wherein Xaa at position 64 is a lysine or a glutamic acid
<220>
<221> misc_feature
<222> (65)..(65)
<223> wherein Xaa at position 65 is an alanine, a proline or a serine
<220>
<221> misc feature
<222> (68)..(68)
<223> wherein Xaa at position 68 is a valine or an alanine
<220>
<221> misc feature
<222>
      (70)..(70)
<223> wherein Xaa at position 70 is a threonine or a glutamic acid
<220>
<221> misc_feature
<222>
      (71)..(71)
<223> wherein Xaa at position 71 is a glutamic acid, a glutamine or a
      lysine
<220>
<221> misc_feature
<222> (72)..(72)
<223> wherein Xaa at position 72 is a leucine or a methionine
<220>
<221> misc feature
<222> (73)..(73)
<223> wherein Xaa at position 73 is a serine, an asparagine or an
      aspartic acid
```

<220>

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<221> misc feature
<222> (74)..(74)
<223> wherein Xaa at position 74 is an alanine, a serine or a proline
<220>
<221> misc_feature
<222> (75)..(75)
<223> wherein Xaa at position 75 is an isoleucine, a leucine or a valine
<220>
<221> misc_feature
<222> (76)..(76)
<223> wherein Xaa at position 76 is a serine or an alanine
<220>
<221> misc feature
<222> (77)..(77)
<223> wherein Xaa at position 77 is a methionine, a valine or an isoleucine
<220>
<221> misc feature
<222> (79)..(79)
<223> wherein Xaa at position 79 is a phenylalanine or a tyrosine
<220>
<221> misc_feature
<222> (80)..(80)
<223> wherein Xaa at position 80 is a leucine, a tyrosine or a phenylalanine
<220>
<221> misc feature
<222> (81)..(81)
<223> wherein Xaa at position 81 is an aspartic acid or an asparagine
<220>
<221> misc_feature
<222> (82)..(82)
<223> wherein Xaa at position 82 is a glutamic acid, an asparagine or an
      aspartic acid
<220>
<221> misc_feature
<222> (83)..(83)
<223> wherein Xaa at position 83 is a glutamine or an asparagine
<220>
<221> misc feature
<222> (84)..(84)
<223> wherein Xaa at position 84 is a glutamic acid, a glutamine, a serine
      or a lysine
<220>
<221> misc_feature
<222>
      (85)..(85)
<223> wherein Xaa at position 85 is an asparagine or a lysine
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<220>

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<221> misc feature
<222> (87)..(87)
<223> wherein Xaa at position 87 is a valine or an isoleucine
<220>
<221> misc feature
<222> (89)..(89)
<223> wherein Xaa at position 89 is a lysine or an arginine
<220>
<221> misc_feature
<222> (90)..(90)
<223> wherein Xaa at position 90 is an asparagine, a lysine or a histidine
<220>
<221> misc feature
<222> (92)..(92)
<223> wherein Xaa at position 92 is a glutamine, a glutamic acid, an
      arginine or a proline
<220>
<221> misc_feature
<222> (93)..(93)
<223> wherein Xaa at position 93 is an aspartic acid, a glutamic acid or
      an asparagine
<220>
<221> misc feature
<222> (95)..(95)
<223> wherein Xaa at position 95 is a valine or a threonine
<220>
<221> misc feature
<222> (97)..(97)
<223> wherein Xaa at position 97 is a glutamic acid, an aspartic acid or
      an arginine
<220>
<221> misc feature
<222> (98)..(98)
<223> wherein Xaa at position 98 is a glycine, an alanine, a serine or
      a glutamic acid
<220>
<221> misc_feature
<222> (100)..(100)
<223> wherein Xaa at position 100 is a glycine or a histidine
<220>
<221> misc_feature
      (102)..(102)
<222>
<223> wherein Xaa at position 102 is a an arginine or a histidine
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Cys Xaa Arg Xaa Xaa Leu Xaa Val Xaa Phe Xaa Asp Xaa Gly Trp Xaa
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Xaa Trp Xaa Xaa Xaa Pro Xaa Gly Xaa Xaa Ala Xaa Tyr Cys Xaa Gly
20 25 30

Xaa Cys Xaa Xaa Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Asn His Ala 35 40 45

Xaa Xaa Gln Xaa Xaa Val Xaa Xaa Xaa Asn Xaa Xaa Xaa Pro Xaa 50 60

Xaa Cys Cys Xaa Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Leu Xaa Xaa 65 70 75 80

Xaa Xaa Xaa Xaa Val Xaa Leu Xaa Xaa Tyr Xaa Xaa Met Xaa Val 85 90 95

Xaa Xaa Cys Xaa Cys Xaa 100

<210> 7

<211> 102

<212> PRT

<213> Xenopus sp.

<220>

<223> Vg1 protein sequence with osteogenic activity

<400> 7

Cys Lys Lys Arg His Leu Tyr Val Glu Phe Lys Asp Val Gly Trp Gln
1 5 10 15

Asn Trp Val Ile Ala Pro Gln Gly Tyr Met Ala Asn Tyr Cys Tyr Gly
20 25 30

Glu Cys Pro Tyr Pro Leu Thr Glu Ile Leu Asn Gly Ser Asn His Ala 35 40 45

Ile Leu Gln Thr Leu Val His Ser Ile Glu Pro Glu Asp Ile Pro Leu 50 55 60

Pro Cys Cys Val Pro Thr Lys Met Ser Pro Ile Ser Met Leu Phe Tyr 65 70 75 80

Asp Asn Asn Asp Asn Val Val Leu Arg His Tyr Glu Asn Met Ala Val 85 90 95

Asp Glu Cys Gly Cys Arg 100

<210> 8

<211> 102

<212> PRT

<213> Drosophila sp.

<220>

<223> DPP protein sequence with osteogenic activity

<400> 8

Cys Arg Arg His Ser Leu Tyr Val Asp Phe Ser Asp Val Gly Trp Asp 1 5 10 15

Asp Trp Ile Val Ala Pro Leu Gly Tyr Asp Ala Tyr Tyr Cys His Gly
20 25 30

Lys Cys Pro Phe Pro Leu Ala Asp His Phe Asn Ser Thr Asn His Ala 35 40 45

Val Val Gln Thr Leu Val Asn Asn Asn Pro Gly Lys Val Pro Lys 50 55 60

Ala Cys Cys Val Pro Thr Gln Leu Asp Ser Val Ala Met Leu Tyr Leu 65 70 75 80

Asn Asp Gln Ser Thr Val Val Leu Lys Asn Tyr Gln Glu Met Thr Val 85 90 95

Val Gly Cys Gly Cys Arg 100

<210> 9

<211> 107

<212> PRT

<213> mammalian

<220>

<223> OP1 protein sequence with osteogenic activity

<400> 9

His Gln Arg Gln Ala Cys Lys Lys His Glu Leu Tyr Val Ser Phe Arg
1 5 10 15

Asp Leu Gly Trp Gln Asp Trp Ile Ile Ala Pro Glu Gly Tyr Ala Ala 20 25 30

Tyr Tyr Cys Glu Gly Glu Cys Ala Phe Pro Leu Asn Ser Tyr Met Asn 35 40 45

Ala Thr Asn His Ala Ile Val Gln Thr Leu Val His Phe Ile Asn Pro 50 55 60

Glu Thr Val Pro Lys Pro Cys Cys Ala Pro Thr Gln Leu Asn Ala Ile 65 70 75 80

Ser Val Leu Tyr Phe Asp Asp Ser Ser Asn Val Ile Leu Lys Lys Tyr
85 90 95

Arg Asn Met Val Val Arg Ala Cys Gly Cys His
100 105

<210> 10 <211> 103

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<212> PRT
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<213> mammalian

<220>

<223> CBP-2a protein sequence with osteogenic activity

<400> 10

Cys Lys Arg His Pro Leu Tyr Val Asp Phe Ser Asp Val Gly Trp Asn 1 5 10 15

Asp Trp Ile Val Ala Pro Pro Gly Tyr His Ala Phe Tyr Cys His Gly
20 25 30

Glu Cys Pro Phe Pro Leu Ala Asp His Leu Asn Ser Thr Asn His Ala 35 40 45

Ile Val Gln Thr Leu Val Asn Ser Val Asn Ser Lys Ile Pro Lys Ala 50 55 60

Cys Cys Val Pro Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu Tyr 65 70 75 80

Leu Asp Glu Asn Glu Lys Val Val Leu Lys Asn Tyr Gln Asp Met Val
85 90 95

Val Glu Gly Cys Gly Cys Arg 100

<210> 11

<211> 100

<212> PRT

<213> mammalian

<220>

<223> CBMP-2b protein sequence with osteogenic activity

<400> 11

Cys Arg Arg His Ser Leu Tyr Val Asp Phe Ser Asp Val Gly Trp Asn
1 5 10 15

Asp Trp Ile Val Ala Pro Pro Gly Tyr Gln Ala Phe Tyr Cys His Gly
20 25 30

Asp Cys Pro Phe Pro Leu Ala Asp His Leu Asn Ser Thr Asn His Ala 35 40 45

Ile Val Gln Thr Leu Val Asn Ser Val Asn Ser Ile Pro Lys Ala Cys 50 55 60

Cys Val Pro Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu Asp Glu 65 70 75 80

Tyr Asp Lys Val Val Leu Lys Asn Tyr Gln Glu Met Val Val Glu Gly
85 90 95

Cys Gly Cys Arg 100 <210> 12 <211> 103 <212> PRT <213> mammalian <220> <223> CBMP-3 protein sequence with osteogenic activity <400> 12 Cys Ala Arg Arg Tyr Leu Lys Val Asp Phe Ala Asp Ile Gly Trp Ser Glu Trp Ile Ile Ser Pro Lys Ser Phe Asp Ala Tyr Tyr Cys Ser Gly Ala Cys Gln Phe Pro Met Pro Lys Ser Leu Lys Pro Ser Asn His Ala 40 Thr Ile Gln Ser Ile Val Arg Ala Val Gly Val Val Pro Gly Ile Pro Glu Pro Cys Cys Val Pro Glu Lys Met Ser Ser Leu Ser Ile Leu Phe Phe Asp Glu Asn Lys Asn Val Val Leu Lys Val Tyr Pro Asn Met Thr Val Glu Ser Cys Ala Cys Arg 100 <210> 13 <211> 98 <212> PRT Artificial Sequence <213> <220> <223> Biosynthetic COP1 protein with osteogenic activity <400> 13 Leu Tyr Val Asp Phe Gln Arg Asp Val Gly Trp Asp Asp Trp Ile Ile Ala Pro Val Asp Phe Asp Ala Tyr Tyr Cys Ser Gly Ala Cys Gln Phe 20 25 30 Pro Ser Ala Asp His Phe Asn Ser Thr Asn His Ala Val Val Gln Thr Leu Val Asn Asn Met Asn Pro Gly Lys Val Pro Lys Pro Cys Cys Val

Pro Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu Asp Glu Asn Ser

65 70 75 80

Thr Val Val Leu Lys Asn Tyr Gln Glu Met Thr Val Val Gly Cys Gly 85 90 95

Cys Arg

<210> 14

<211> 98

<212> PRT

<213> Artificial Sequence

<220>

<223> Biosynthetic COP3 protein with osteogenic activity

<400> 14

Leu Tyr Val Asp Phe Gln Arg Asp Val Gly Trp Asp Asp Trp Ile Val

1 10 15

Ala Pro Pro Gly Tyr Gln Ala Phe Tyr Cys Ser Gly Ala Cys Gln Phe 20 25 30

Pro Ser Ala Asp His Phe Asn Ser Thr Asn His Ala Val Gln Thr
35 40 45

Leu Val Asn Asn Met Asn Pro Gly Lys Val Pro Lys Pro Cys Cys Val
50 60

Pro Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu Asp Glu Asn Glu 65 70 75 80

Lys Val Val Leu Lys Asn Tyr Gln Glu Met Val Val Glu Gly Cys Gly
85 90 95

Cys Arg

<210> 15

<211> 97

<212> PRT

<213> Artificial Sequence

<220>

<223> Biosynthetic COP4 protein with osteogenic activity

<400> 15

Leu Tyr Val Asp Phe Ser Asp Val Gly Trp Asp Asp Trp Ile Val Ala 1 5 10 15

Pro Pro Gly Tyr Gln Ala Phe Tyr Cys Ser Gly Ala Cys Gln Phe Pro 20 25 30

Ser Ala Asp His Phe Asn Ser Thr Asn His Ala Val Val Gln Thr Leu 35 40 45

Val Asn Asn Met Asn Pro Gly Lys Val Pro Lys Pro Cys Cys Val Pro Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu Asp Glu Asn Glu Lys Val Val Leu Lys Asn Tyr Gln Glu Met Val Val Glu Gly Cys Gly Cys Arg <210> 16 <211> 97 <212> PRT <213> Artificial Sequence <220> <223> Biosynhetic COP16 protein with osteogenic activity <400> 16 Leu Tyr Val Asp Phe Ser Asp Val Gly Trp Asp Asp Trp Ile Val Ala Pro Pro Gly Tyr Gln Ala Phe Tyr Cys Ser Gly Ala Cys Gln Phe Pro Ser Ala Asp His Phe Asn Ser Thr Asn His Ala Val Val Gln Thr Leu Val Asn Asn Met Asn Pro Gly Lys Val Pro Lys Pro Cys Cys Val Pro 50 Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu Asp Glu Asn Glu Lys Val Val Leu Lys Asn Tyr Gln Glu Met Val Val Glu Gly Cys Gly Cys Arg <210> 17 <211> 17 <212> PRT <213> Artificial Sequence <220> <223> peptide fragment

<400> 17

Ser Phe Asp Ala Tyr Tyr Cys Ser Gly Ala Cys Gln Phe Pro Met Pro

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<210> 18
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<212> PRT
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<223> peptide fragment
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Ser Leu Lys Pro Ser Asn Tyr Ala Thr Ile Gln Ser Ile Val
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<210> 19
<211> 21
<212> PRT
<213> Artificial Sequence
<220>
<223> peptide fragment
<400> 19
Ala Cys Cys Val Pro Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu
                                   10
Asp Glu Asn Glu Lys
<210> 20
<211> 13
<212> PRT
<213> Artificial Sequence
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<223> peptide fragment
<400> 20
Met Ser Ser Leu Ser Ile Leu Phe Phe Asp Glu Asn Lys
<210> 21
<211> 10
<212> PRT
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<223> peptide fragment
<400> 21
Ser Gln Glu Leu Tyr Val Asp Phe Gln Arg
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Lys

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<211> 11
<212> PRT
<213> Artificial Sequence
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<223> peptide fragment
<400> 22
Phe Leu His Cys Gln Phe Ser Glu Arg Asn Ser
<210> 23
<211> 15
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Thr Val Gly Gln Leu Asn Glu Gln Ser Ser Glu Pro Asn Ile Tyr
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Leu Tyr Asp Pro Met Val Val
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Val Gly Val Val Pro Gly Ile Pro Glu Pro Cys Cys Val Pro Glu
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<220>
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Val Asp Phe Ala Asp Ile Gly
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Val Pro Lys Pro Cys Cys Ala Pro Thr
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Ile Asn Ile Ala Asn Tyr Leu
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<211> 13
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Asp Asn His Val Leu Thr Met Phe Pro Ile Ala Ile Asn
<210> 30
<211> 16
<212> PRT
<213> Artificial Sequence
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<223> peptide fragment
<220>
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<221> misc_feature
<222> (15)..(15)
<223> wherein Xaa at position 15 is any amino acid
<400> 30
Asp Glu Gln Thr Leu Lys Lys Ala Arg Arg Lys Gln Trp Ile Xaa Pro
<210> 31
<211> 11
<212> PRT
<213> Artificial Sequence
<220>
<223> peptide fragment
<220>
<221> misc feature
<222> (4)..(4)
<223> wherein Xaa at position 4 is any.amino acid
<220>
<221> misc_feature
<222> (10)..(10)
<223> wherein Xaa at position 10 is any amino acid
<400> 31
Asp Ile Gly Xaa Ser Glu Trp Ile Ile Xaa Pro
<210> 32
<211> 17
<212> PRT
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<220>
<223> peptide fragment
<220>
<221> misc_feature
<222>
      (15)..(16)
<223> wherein Xaa at positions 15 and 16 is any amino acid
<400> 32
Ser Ile Val Arg Ala Val Gly Val Pro Gly Ile Pro Glu Pro Xaa Xaa
                5
                                    10
Val
<210> 33
<211> 13
<212> PRT
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<220>
<223> peptide fragment
<220>
<221> misc_feature
<222>
      (2)..(2)
<223> wherein Xaa at position 2 is any amino acid
<400> 33
Asp Xaa Ile Val Ala Pro Pro Gln Tyr His Ala Phe Tyr
<210> 34
<211> 17
<212> PRT
<213> Artificial Sequence
<220>
<223> peptide fragment
<400> 34
Asp Glu Asn Lys Asn Val Val Leu Lys Val Tyr Pro Asn Met Thr Val
                                    10
Glu
<210> 35
<211> 18
<212> PRT
<213> Artificial Sequence
<220>
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<220>
<221> misc_feature
<222> (13)..(13)
<223> wherein Xaa at position 13 is any amino acid
<220>
<221> misc_feature
<222>
      (16)..(16)
<223> wherein Xaa at position 16 is any amino acid
<400> 35
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Lys Gln
<210> 36
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<212> PRT
<213> Artificial Sequence
<220>
<223> peptide fragment
<220>
<221> misc_feature
<222>
      (19)..(19)
<223> wherein Xaa at position 19 is any amino acid
<400> 36
Asp Glu Gln Thr Leu Lys Lys Ala Arg Arg Lys Gln Trp Ile Glu Pro
                5
                                   10
Arg Asn Xaa Ala Arg Arg Tyr Leu
            20
<210> 37
<211> 20
<212> PRT
<213> Artificial Sequence
<220>
<223> peptide fragment
<220>
<221> misc_feature
<222> (12)..(12)
<223> wherein Xaa at position 12 is any amino acid
<220>
<221> misc feature
<222> (14)..(14)
<223> wherein Xaa at position 14 is any amino acid
<220>
<221> misc feature
<222> (17)..(18)
<223> wherein Xaa at positions 17-18 is any amino acid
<400> 37
Ala Arg Arg Lys Gln Trp Ile Glu Pro Arg Asn Xaa Ala Xaa Arg Tyr
                                    10
Xaa Xaa Val Asp
            20
<210> 38
<211> 23
<212> PRT
<213> Artificial Sequence
<220>
<223> peptide fragment
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<220>
<221> misc feature
<222>
      (2)..(2)
<223> wherein Xaa at position 2 is any amino acid
<220>
<221> misc feature
<222>
      (8)..(8)
<223> wherein Xaa at position 8 is any amino acid
<220>
<221> misc_feature
<222> (10)..(10)
<223> wherein Xaa at position 10 is any amino acid
<220>
<221> misc feature
<222> (12)..(13)
<223> wherein Xaa at positions 12-13 is any amino acid
<220>
<221> misc feature
<222> (19)..(19)
<223> wherein Xaa at position 19 is any amino acid
<220>
<221> misc_feature
<222>
      (21)..(22)
<223>
      wherein Xaa at positions 21-22 is any amino acid
<400> 38
Arg Xaa Gln Trp Ile Glu Pro Xaa Asn Xaa Ala Xaa Xaa Tyr Leu Lys
                                    10
Val Asp Xaa Ala Xaa Xaa Gly
            20
<210> 39
<211> 97
<212> PRT
<213> mammalian
<220>
<223> OP1 shorter sequence capable of inducing endochondral bone formation
<400> 39
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Pro Glu Gly Tyr Ala Ala Tyr Tyr Cys Glu Gly Glu Cys Ala Phe Pro
            20
Leu Asn Ser Tyr Met Asn Ala Thr Asn His Ala Ile Val Gln Thr Leu
        35
                            40
                                                45
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Val His Phe Ile Asn Pro Glu Thr Val Pro Lys Pro Cys Cys Ala Pro Thr Gln Leu Asn Ala Ile Ser Val Leu Tyr Phe Asp Asp Ser Ser Asn 65 70 Val Ile Leu Lys Lys Tyr Arg Asn Met Val Val Arg Ala Cys Gly Cys His <210> 40 4805 <211> <212> DNA <213> mammalian <220> <223> genomic sequence of OP1 <220> <221> misc feature <223> approximately 1000 bases are missing between position 1883 and 188 <400> 40 ggaggtatag gagctctctt cgattttagc aaaccaggag tccgaagatc taaqqaqaqc 60 tgggggtttg actocgagag ctogagoagt coccaagaco tggtottgac toacqaqtta 120 gactccactc agaggctgac tgtctccagg gtctacacct ctaagggcga cactgggctc 180 aagcagactg ccgttttcta tatgggatga gccttcacag ggcagccagt tqqqatqqqt 240 tgaggtttgg ctgtagacat cagaaaccca agtcaaatgc gcttcaacca gtagaaaatt 300 caccagcccg cagagctaag gttgggtgga cattagggtt ggttgatcca qqaqctcaac 360 420 tcaaggccac agctgtagtt ggccaggggg gcttcattat tttttgctcc tggqcaqtaq 480 gaggaagaga atgaatgtct ctccatgggt ctttcttagg aatgtgggaa ctttttccag 540 aagtetetat gtettttagt ttgtgttggg teaettgeee tteetgaace aetteetgae 600 tcctggacag gatgtgcact gatgagctta gctttgggga tctaatagtg actttacaaa 660 gcctctttga gaaggtgaca ttggaaccaa ggcttgagca gacacaacaa agattgcagg 720 gaggggcatt gcaggtggag gaaacggcac atgcaagagc cctgcgtggg agtgagcttq 780 gtgtttggtc aatcagttgt cagagcacac cgggccctgt cagcaggcac agcctgggcc 840

900

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gac tgg Asp Tr															96
gcc tgo Ala Cys															144
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Ala Cys Gln Phe Pro Ser Ala Asp His Phe Asn Ser Thr Asn His Ala 35 40 45

Val Val Gln Thr Leu Val Asn Asn Met Asn Pro Gly Lys Val Pro Lys 50 55 60

Pro Cys Cys Val Pro Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu 65 70 75 80

Asp Glu Asn Ser Thr Val Val Leu Lys Asn Tyr Gln Glu Met Thr Val 85 90 95

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Asp Trp Ile Ile Ala Pro Ser Gly Tyr His Ala Asn Tyr Cys Glu Gly
20 25 30

Glu Cys Pro Ser His Ile Ala Gly Thr Ser Gly Ser Ser Leu Ser Phe

35 40 45

His Ser Thr Val Ile Asn His Tyr Arg Met Arg Gly His Ser Pro Phe 50 60

Ala Asn Leu Lys Ser Cys Cys Val Pro Thr Lys Leu Arg Pro Met Ser 65 70 75 80

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Trp Ile Ile Ala Pro Thr Gly Tyr Tyr Gly Asn Tyr Cys Glu Gly Ser 20 25 30

Cys Pro Ala Tyr Leu Ala Gly Val Pro Gly Ser Ala Ser Ser Phe His 35 40 45

Thr Ala Val Val Asn Gln Tyr Arg Met Arg Gly Leu Asn Pro Gly Thr
50 55 60

Lys Val Asn Ser Cys Cys Ile Pro Thr Lys Leu Ser Thr Met Ser Met 65 70 75 80

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Pro Cys Pro Tyr Ile Trp Ser Leu Leu Asp Thr Gln Tyr Ser Lys Val 35 40 45

Leu Ala Leu Tyr Asn Gln His Asn Pro Gly Ala Ser Ala Ala Pro Cys
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20 25 30

Ala Cys Pro Tyr Leu Trp Ser Leu Ser Asp Thr Gln His Ser Arg Val 35 40 45

Leu Ser Leu Tyr Asn Thr Ile Asn Pro Glu Ala Ser Ala Ser Pro Cys 50 55 60

Cys Val Ser Gln Asp Leu Glu Pro Leu Thr Ile Leu Tyr Tyr Ile Gly 65 70 75 80

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Pro Cys Pro Tyr Leu Arg Ser Leu Ala Asp Thr Thr His Ser Thr Val

Leu Gly Leu Tyr Asn Thr Leu Asn Pro Glu Ala Ser Ala Ser Pro Cys 50 55 60

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Trp Pro Gln Ser Asp Arg Asn Pro Arg Tyr Gly Asn His Val Leu 35 40 45

Leu Leu Lys Met Gln Ala Arg Gly Ala Ala Leu Ala Arg Pro Pro Cys 50 55 60

Cys Val Pro Thr Ala Tyr Ala Gly Lys Leu Leu Ile Ser Leu Ser Glu 65 70 75 80

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                            40
Ala Pro Pro Thr Pro Ala Gln Pro Tyr Ser Leu Leu Pro Gly Ala Gln
    50
Pro Cys Cys Ala Ala Leu Pro Gly Thr Met Arg Pro Leu His Val Arg
Thr Thr Ser Asp Gly Gly Tyr Ser Phe Lys Tyr Glu Xaa Asn Leu Leu
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Thr Gln His Cys Ala Cys Ile
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tet egt etg gat etg gae gtt egt ace gae cae aaa gae etg tet gat
                                                                      96
Ser Arg Leu Asp Leu Asp Val Arg Thr Asp His Lys Asp Leu Ser Asp
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cac ctg gtt ctg gtc gac ctg gct cgt aac gac ctg gct cgt atc gtt
                                                                     144
His Leu Val Leu Val Asp Leu Ala Arg Asn Asp Leu Ala Arg Ile Val
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					cag Gln 70											240
					gag Glu											288
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					gaa Glu											432
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					gca Ala											576
					tgg Trp											624
					cat His											672
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					aag Lys											768
atc Ile	agc Ser	atg Met	ctg Leu 260	tac Tyr	ctg Leu	gac Asp	gag Glu	aat Asn 265	gag Glu	aag Lys	gtg Val	gtg Val	ctg Leu 270	aag Lys	aac Asn	816
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165

Tyr Gln Glu Met Val Val Glu Gly Cys Gly Cys Arg

280

275

175

170

Leu Asn Asp Ala Gln Ala Pro Lys Asp Pro Asn Gly Leu Tyr Val Asp 180 185 Phe Ser Asp Val Gly Trp Asp Asp Trp Ile Val Ala Pro Pro Gly Tyr 195 200 Gln Ala Phe Tyr Cys His Gly Glu Cys Pro Phe Pro Leu Ala Asp His 210 215 220 Phe Asn Ser Thr Asn His Ala Val Val Gln Thr Leu Val Asn Ser Val 225 230 Asn Ser Lys Ile Pro Lys Ala Cys Cys Val Pro Thr Glu Leu Ser Ala 245 250 Ile Ser Met Leu Tyr Leu Asp Glu Asn Glu Lys Val Val Leu Lys Asn 260 265 Tyr Gln Glu Met Val Val Glu Gly Cys Gly Cys Arg 275 280 <210> 55 <211> 15 <212> PRT <213> Artificial Sequence <220> <223> BOP <400> 55 Ser Phe Asp Ala Tyr Tyr Cys Ser Gly Ala Cys Gln Phe Pro Ser 5 10 <210> 56 <211> 15 <212> PRT <213> Artificial Sequence <220> <223> DPP <400> 56 Gly Tyr Asp Ala Tyr Tyr Cys His Gly Lys Cys Pro Phe Phe Leu 10 5 <210> 57

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